

PHASE I HCV TRIAL: STUDY PROTOCOL																													
WEEK:	0	IV+ORAL TREATMENT:										ORAL TREATMENT:										FOLLOW UP:							
ORAL TREATMENT		1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	24	28	32	36	40			
		X	X	X	X	X	X	X	X	X	X	X	X	X	X	X	X	X	X	X	X								
		2	2	2	2	2	2	2	2	2	2																		
PHYSICIAN VISIT	X	X	X	X	X	X	X	X	X	X	X		X		X		X		X		X	X	X	X	X	X			
SMA (6)	X				X			X					X				X				X	X			X				
CBC	X				X			X					X				X				X	X			X				
HCV-PCR RNA	X				X			X					X				X				X	X			X				
QUANTITATIVE													X				X				X				X				
ANTI HCV	X																												
ANTI HCV	X																												
CONFIRMATION																													
HCV-PCR	X																												
GENOTYPE																													
CRYOGLOBULINS	X																												
LIVER BIOPSY	X																												
ABDOMINAL US	X																												
PT, PTT	X				X			X					X				X				X	X			X				
AFP	X																												
NH3	X																												
ESR	X																												
ANF	X																												
C3	X																												
ANTI	X																												
MITOCHONDRIA																													
ANTI SMOOTH	X																												
MUSCLE																													
ANTI Parietal	X																												
CELL																													
IEP	X																												
HIV	X																												
HBSAG	X																												

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FIGURE 2

SCORING & GRADING SYSTEMS

Table 1. HAI for numerical scoring of liver biopsy specimens

I. Periportal +/- Bridging Necrosis	Score	II. Intralobular Degeneration* and Focal Necrosis	Score	III. Portal Inflammation	Score	IV. Fibrosis	Score
A. None	0	A. None	0	A. No portal inflammation	0	A. No fibrosis	0
B. Mild piecemeal necrosis	1	B. Mild (acidophilic bodies, ballooning degeneration and/or scattered foci of hepatocellular necrosis in <1/3 of lobules or nodules)	1	B. Mild (sprinkling of inflammatory cells in <1/3 of portal tracts)	1	B. Fibrous portal expansion	1
C. Moderate piecemeal necrosis (involves less than 50% of the circumference of most portal tracts)	3	C. Moderate (involvement of 1/3-2/3 of lobules or nodules)	3	C. Moderate (increased inflammatory cells in 1/3- 2/3 of portal tracts)	3	C. Bridging fibrosis (portal-portal or portal- central linkage)	3
D. Marked piecemeal necrosis (involves more than 50% of the circumference of most portal tracts)	4	D. Marked (involvement of >2/3 of lobules or nodules)	4	D. Marked (dense packing of inflammatory cells in >2/3 of portal tracts)	4	D. Cirrhosis†	4
E. Moderate piecemeal necrosis <i>plus</i> bridging necrosis‡	5						
F. Marked piecemeal necrosis <i>plus</i> bridging necrosis‡	6						
G. Multilobular necrosis§	10						

NOTE. HAI score is the combined scores for necrosis, inflammation, and fibrosis.

*Degeneration—acidophil bodies, ballooning; focal necrosis—scattered foci of hepatocellular necrosis.

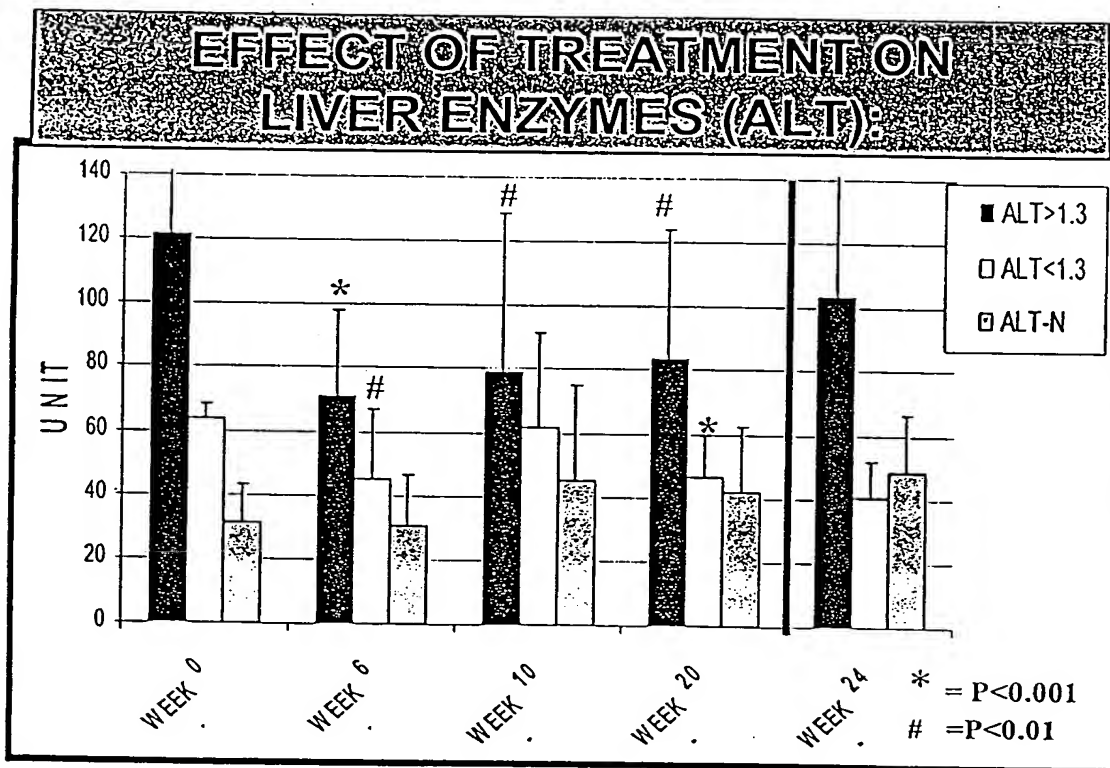
†Loss of normal hepatic lobular architecture with fibrous septae separating and surrounding nodules.

‡Bridging is defined as ≥ 2 bridges in the liver biopsy specimen; no distinction is made between portal-portal and portal-central linkage.

§Two or more contiguous lobules with panlobular necrosis.

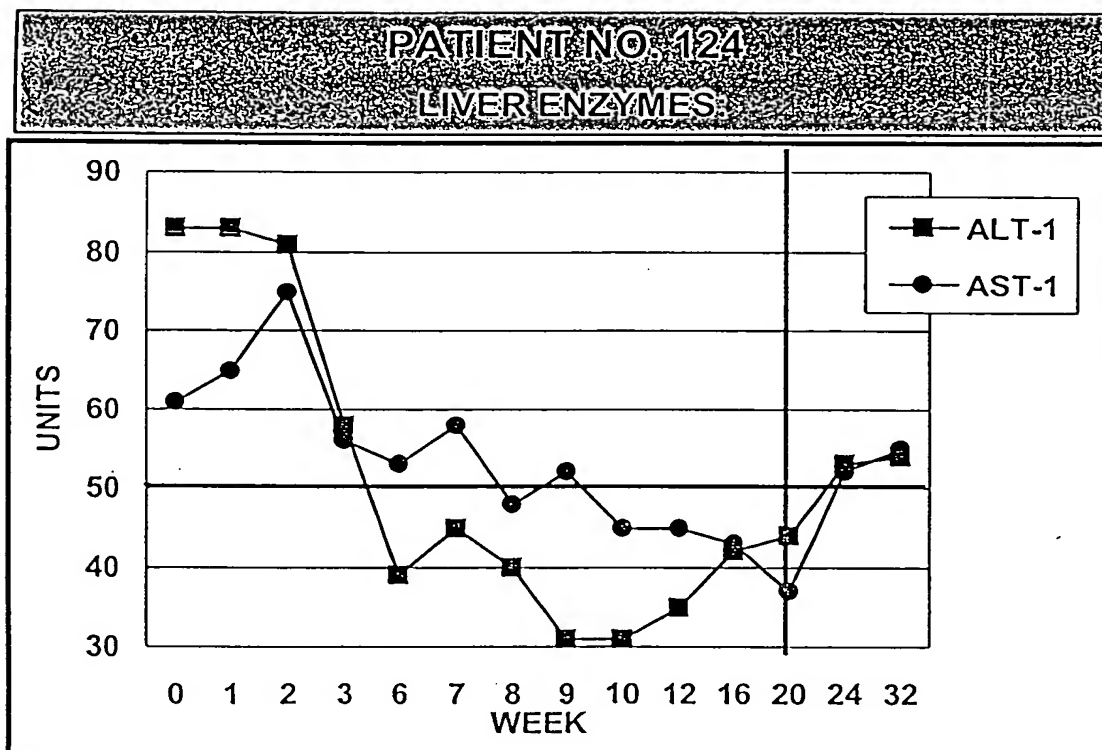
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FIGURE 3



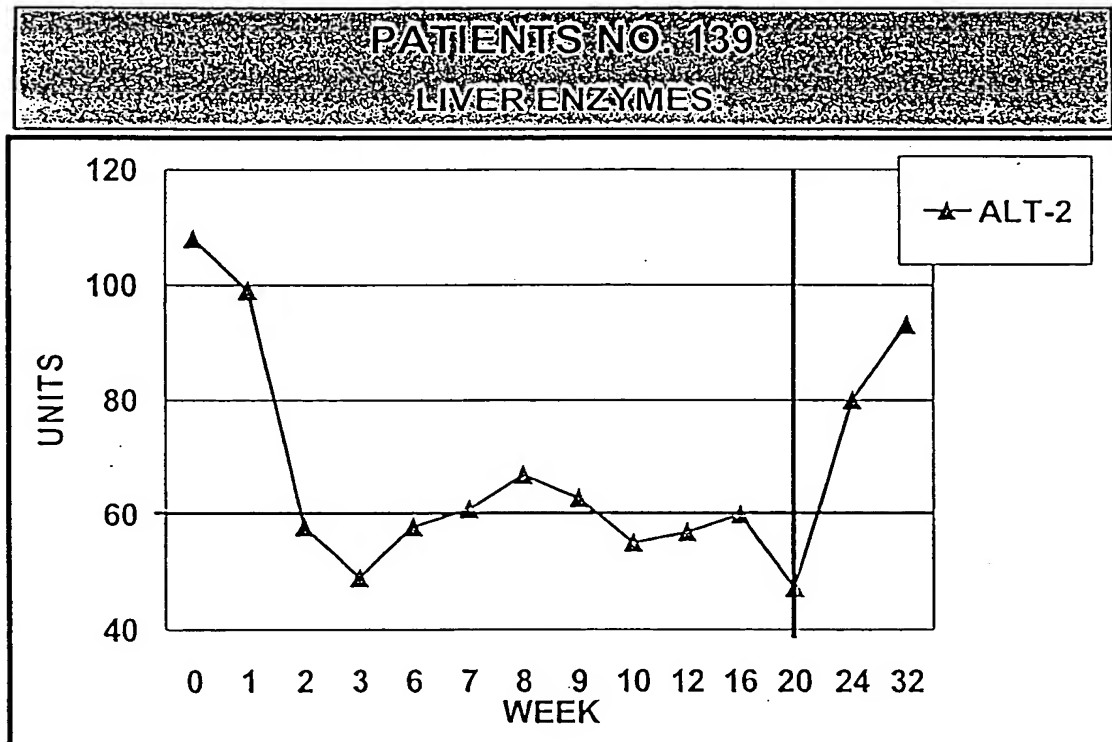
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FIGURE 4



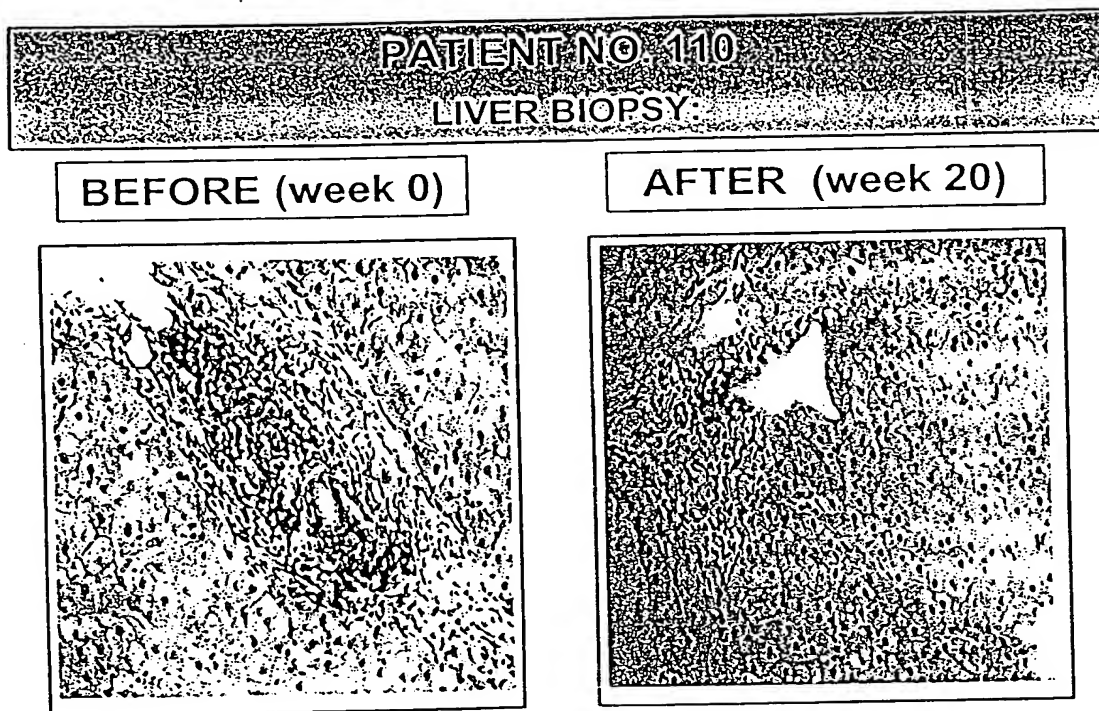
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FIGURE 5



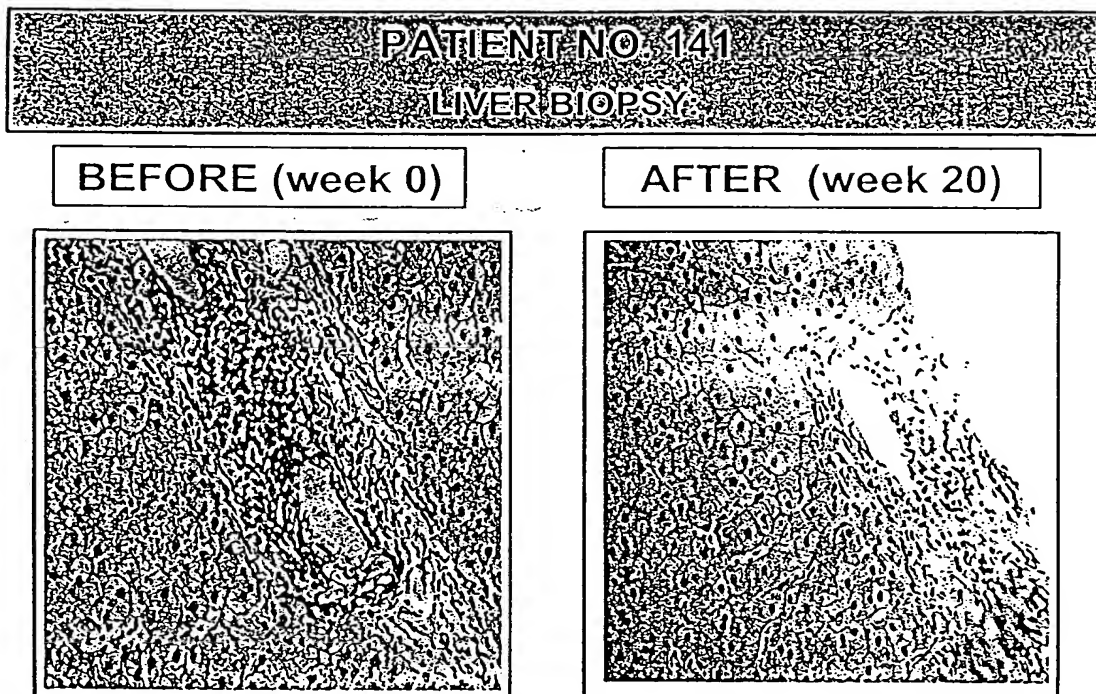
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FIGURE 6



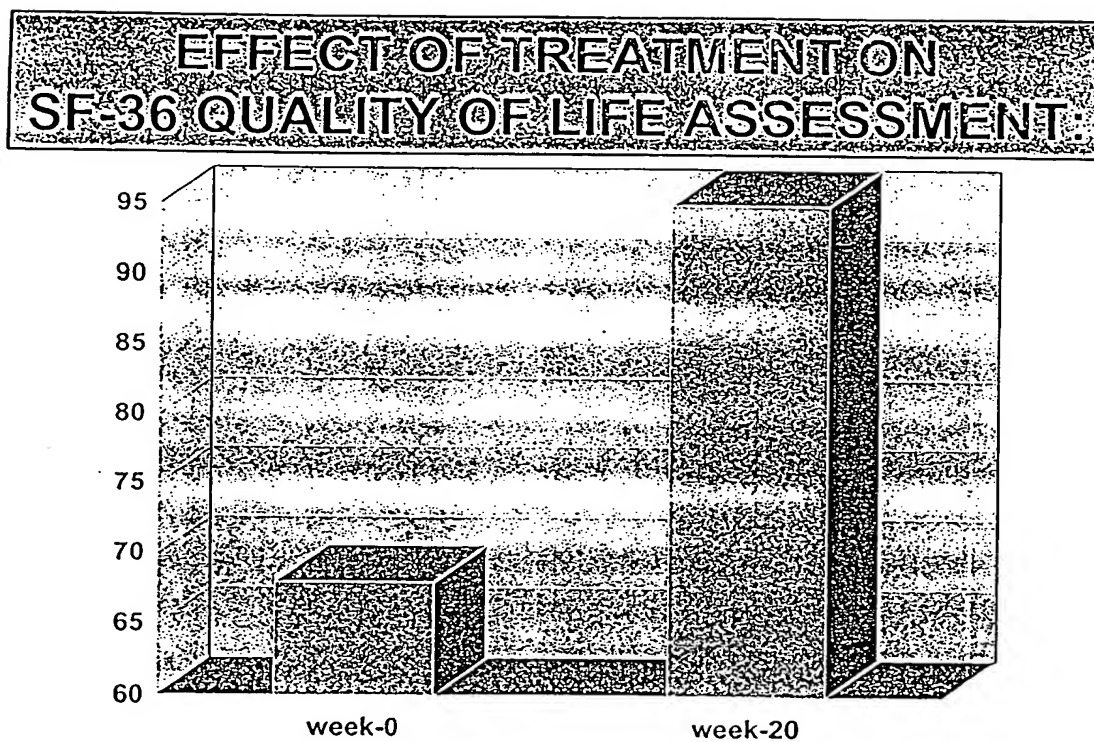
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FIGURE 7



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FIGURE 8



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